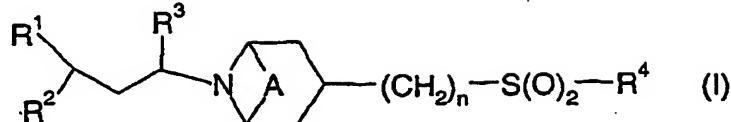


CLAIMS

1. A compound of formula (I):



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wherein:

A is absent or is $(CH_2)_2$;

10 R¹ is $C(O)NR^{10}R^{11}$, $C(O)_2R^{12}$, $NR^{13}C(O)R^{14}$, $NR^{15}C(O)NR^{16}R^{17}$, $NR^{18}C(O)_2R^{19}$, heterocyclyl, aryl or heteroaryl;

15 R¹⁰, R¹³, R¹⁵, R¹⁶ and R¹⁸ are hydrogen or C₁₋₆ alkyl;

20 R¹¹, R¹², R¹⁴, R¹⁷ and R¹⁹ are C₁₋₈ alkyl (optionally substituted by halo, hydroxy, C₁₋₆

alkoxy, C₁₋₆ haloalkoxy, C₃₋₆ cycloalkyl (optionally substituted by halo), C₅₋₆

cycloalkenyl, S(C₁₋₄ alkyl), S(O)(C₁₋₄ alkyl), S(O)₂(C₁₋₄ alkyl), heteroaryl, aryl,

heteroaryloxy or aryloxy), aryl, heteroaryl, C₃₋₇ cycloalkyl (optionally substituted by

halo or C₁₋₄ alkyl), C₄₋₇ cycloalkyl fused to a phenyl ring, C₅₋₇ cycloalkenyl, or,

25 heterocyclyl (itself optionally substituted by oxo, C(O)(C₁₋₆ alkyl), S(O)₂(C₁₋₆ alkyl),

halo or C₁₋₄ alkyl); or R¹¹, R¹², R¹⁴ and R¹⁷ can also be hydrogen;

or R¹⁰ and R¹¹, and/or R¹⁶ and R¹⁷ may join to form a 4-, 5- or 6-membered ring which

optionally includes a nitrogen, oxygen or sulphur atom, said ring being optionally

substituted by C₁₋₆ alkyl, S(O)₂(C₁₋₆ alkyl) or C(O)(C₁₋₆ alkyl);

20 R² is phenyl, heteroaryl or C₃₋₇ cycloalkyl;

R³ is H or C₁₋₄ alkyl;

R⁴ is heterocyclyl;

n is 1, 2 or 3;

30 aryl, phenyl and heteroaryl moieties are independently optionally substituted by one or

more of halo, cyano, nitro, hydroxy, $OC(O)NR^{20}R^{21}$, $NR^{22}R^{23}$, $NR^{24}C(O)R^{25}$,

$NR^{26}C(O)NR^{27}R^{28}$, $S(O)_2NR^{29}R^{30}$, $NR^{31}S(O)_2R^{32}$, $C(O)NR^{33}R^{34}$, CO_2R^{36} ,

$NR^{37}CO_2R^{38}$, $S(O)_4R^{39}$, $OS(O)_2R^{40}$, C₁₋₆ alkyl (optionally mono-substituted by

$S(O)_2R^{50}$ or $C(O)NR^{51}R^{52}$), C₂₋₆ alkenyl, C₂₋₆ alkynyl, C₃₋₁₀ cycloalkyl, C₁₋₆ haloalkyl,

C₁₋₆ alkoxy(C₁₋₆)alkyl, C₁₋₆ alkoxy, C₁₋₆ haloalkoxy, phenyl, phenyl(C₁₋₄)alkyl,

phenoxy, phenylthio, phenylS(O), phenylS(O)₂, phenyl(C₁₋₄)alkoxy, heteroaryl,

heteroaryl(C₁₋₄)alkyl, heteroaryloxy or heteroaryl(C₁₋₄)alkoxy; wherein any of the immediately foregoing phenyl and heteroaryl moieties are optionally substituted with halo, hydroxy, nitro, S(C₁₋₄ alkyl), S(O)(C₁₋₄ alkyl), S(O)₂(C₁₋₄ alkyl), S(O)₂NH₂, S(O)₂NH(C₁₋₄ alkyl), S(O)₂N(C₁₋₄ alkyl)₂, cyano, C₁₋₄ alkyl, C₁₋₄ alkoxy, C(O)NH₂, C(O)NH(C₁₋₄ alkyl), C(O)N(C₁₋₄ alkyl)₂, CO₂H, CO₂(C₁₋₄ alkyl), NHC(O)(C₁₋₄ alkyl), NHS(O)₂(C₁₋₄ alkyl), CF₃ or OCF₃;

unless otherwise stated heterocycl is optionally substituted by C₁₋₆ alkyl [optionally substituted by phenyl { which itself optionally substituted by halo, C₁₋₄ alkyl, C₁₋₄ alkoxy, cyano, nitro, CF₃, OCF₃, (C₁₋₄ alkyl)C(O)NH, S(O)₂NH₂, C₁₋₄ alkylthio,

10 S(O)(C₁₋₄ alkyl) or S(O)₂(C₁₋₄ alkyl)} or heteroaryl {which itself optionally substituted by halo, C₁₋₄ alkyl, C₁₋₄ alkoxy, cyano, nitro, CF₃, (C₁₋₄ alkyl)C(O)NH, S(O)₂NH₂, C₁₋₄ alkylthio, S(O)(C₁₋₄ alkyl) or S(O)₂(C₁₋₄ alkyl)}], phenyl {optionally substituted by halo, C₁₋₄ alkyl, C₁₋₄ alkoxy, cyano, nitro, CF₃, OCF₃, (C₁₋₄ alkyl)C(O)NH, S(O)₂NH₂, C₁₋₄ alkylthio, S(O)(C₁₋₄ alkyl) or S(O)₂(C₁₋₄ alkyl)}, heteroaryl {optionally substituted by halo, C₁₋₄ alkyl, C₁₋₄ alkoxy, cyano, nitro, CF₃, (C₁₋₄ alkyl)C(O)NH, S(O)₂NH₂, C₁₋₄ alkylthio, S(O)(C₁₋₄ alkyl) or S(O)₂(C₁₋₄ alkyl)}, S(O)₂NR⁴⁰R⁴¹, C(O)R⁴², C(O)₂(C₁₋₆ alkyl) (such as tert-butoxycarbonyl), C(O)₂(phenyl(C₁₋₂ alkyl)) (such as benzyloxycarbonyl), C(O)NHR⁴³, S(O)₂R⁴⁴, NHS(O)₂NHR⁴⁵, NHC(O)R⁴⁶, NHC(O)NHR⁴⁷ or NHS(O)₂R⁴⁸, provided none of these last four substituents is linked to a ring nitrogen;

k, l and q are, independently, 0, 1 or 2;

R^{20} , R^{22} , R^{24} , R^{26} , R^{27} , R^{29} , R^{31} , R^{33} , R^{37} , R^{40} and R^{51} are, independently, hydrogen or C_{1-6} alkyl;

$R^{21}, R^{23}, R^{25}, R^{28}, R^{30}, R^{32}, R^{34}, R^{36}, R^{38}, R^{39}, R^{41}, R^{42}, R^{43}, R^{44}, R^{45}, R^{46}, R^{47}, R^{48}, R^{49}, R^{50}$ and R^{52} are, independently, C_{1-6} alkyl (optionally substituted by halo, hydroxy, C_{1-6} alkoxy, C_{1-6} haloalkoxy, C_{3-6} cycloalkyl, C_{5-6} cycloalkenyl, $S(C_{1-4}$ alkyl), $S(O)(C_{1-4}$ alkyl), $S(O)_2(C_{1-4}$ alkyl), heteroaryl, phenyl, heteroaryloxy or phenoxy), C_{3-7} cycloalkyl, phenyl or heteroaryl; wherein any of the immediately foregoing phenyl and heteroaryl moieties are optionally substituted with halo, hydroxy, nitro, $S(C_{1-4}$ alkyl),

30 S(O)(C₁₋₄ alkyl), S(O)₂(C₁₋₄ alkyl), S(O)₂NH₂, S(O)₂NH(C₁₋₄ alkyl), S(O)₂N(C₁₋₄ alkyl)₂, cyano, C₁₋₄ alkyl, C₁₋₄ alkoxy, C(O)NH₂, C(O)NH(C₁₋₄ alkyl), C(O)N(C₁₋₄ alkyl)₂, CO₂H, CO₂(C₁₋₄ alkyl), NHC(O)(C₁₋₄ alkyl), NHS(O)₂(C₁₋₄ alkyl), C(O)(C₁₋₄ alkyl), CF₃ or OCF₃;

R^{21} , R^{23} , R^{25} , R^{28} , R^{30} , R^{34} , R^{35} , R^{36} , R^{41} , R^{42} , R^{43} , R^{45} , R^{46} , R^{47} and R^{52} may additionally be hydrogen;
or a pharmaceutically acceptable salt thereof or a solvate thereof.

5 2. A compound as claimed in claim 1 wherein R^1 is $NR^{13}C(O)R^{14}$, wherein R^{13} and R^{14} are as defined in claim 1.

3. A compound as claimed in claim 1 or 2 wherein R^1 is optionally substituted aryl or 10 optionally substituted heteroaryl, wherein the optional substituents are as recited in claim 1.

4. A compound as claimed in claim 1, 2 or 3 wherein R^1 is optionally substituted heterocyclyl.

15 5. A compound as claimed in any one of the preceding claims wherein R^2 is phenyl optionally substituted by halo or CF_3 .

6. A compound as claimed in any one of the preceding claims wherein R^3 is hydrogen.

20 7. A compound as claimed in any one of the preceding claims wherein R^4 is heterocyclyl optionally substituted by oxo, halogen, cyano, hydroxy, C_{1-6} alkyl (itself optionally substituted by halogen, hydroxy, cyano or C_{1-4} alkoxy), C_{2-4} alkenyl, $CO_2(C_{1-4}$ alkyl), $S(O)_2(C_{1-4}$ alkyl), $CH(O)$, $S(O)_2(C_{1-4}$ haloalkyl), $C(O)(C_{1-4}$ alkyl), $C(O)(C_{3-6}$ cycloalkyl), $N(C_{1-4}$ alkyl)₂, $C(O)NH_2$, $C(O)N(C_{1-4}$ alkyl)₂ or $NHC(O)(C_{1-4}$ alkyl).

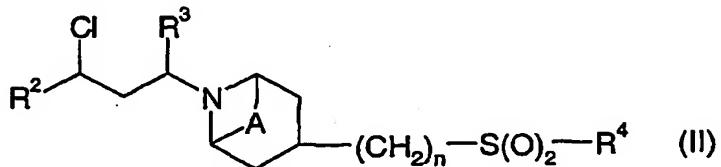
25 8. A compound as claimed in any one of the preceding claims wherein heterocyclyl is piperidinyl, homopiperazinyl, thiomorpholinyl, pyrrolidinyl, piperazinyl, 1,2,3,6-tetrahydropyridinyl, morpholinyl, 2,5-dihydropyrrolyl, azetidinyl, 1,4-oxepanyl, 3-azabicyclo[3.2.1]octan-3-yl, 8-azaspiro[4.5]decanyl or 3-azabicyclo[3.1.0]hex-3-yl.

30 9. A compound as claimed in any one of the preceding claims wherein A is absent.

10. A compound as claimed in any one of the preceding claims wherein n is 2.

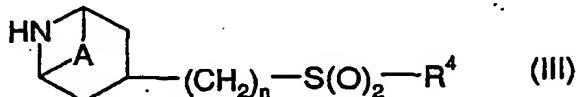
11. A process for preparing a compound as claimed in claim 1, the process comprising:

- when R^1 is an N-linked optionally substituted heterocycle, reacting a compound of formula (II):

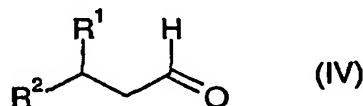


wherein R^2 , R^3 , R^4 , n , A and X are as defined in claim 1, with a compound R^1H (wherein the H is on a heterocycle ring nitrogen atom) wherein R^1 is as defined in claim 1, in the presence of a suitable base and in a suitable solvent;

- when R^3 is hydrogen, coupling a compound of formula (III):



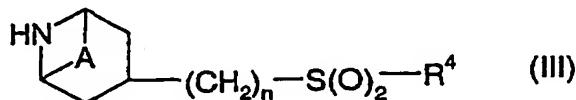
wherein R^4 , n , A and X are as defined in claim 1, with a compound of formula (IV):



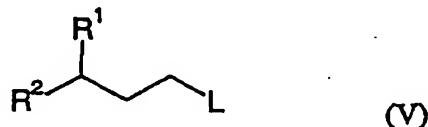
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wherein R^1 and R^2 are as defined in claim 1, in the presence of $NaBH(OAc)_3$ (wherein Ac is $C(O)CH_3$) in a suitable solvent at room temperature; or,

- when R^3 is hydrogen, coupling a compound of formula (III):



wherein R^4 , n , A and X are as defined in claim 1, with a compound of formula (V):



wherein R^1 and R^2 are as defined in claim 1 and L is an activated leaving group, in the presence of a base, in a suitable solvent at a temperature from $60^\circ C$ up to the boiling point of the solvent.

12. A pharmaceutical composition which comprises a compound as claimed in claim 1, or a pharmaceutically acceptable salt thereof or solvate thereof, and a pharmaceutically acceptable adjuvant, diluent or carrier.

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13. A compound as claimed in claim 1, or a pharmaceutically acceptable salt thereof or solvate thereof, for use as a medicament.

14. A compound as claimed in claim 1, or a pharmaceutically acceptable salt thereof or solvate thereof, in the manufacture of a medicament for use in therapy.

15. A method of treating a CCR5 mediated disease state comprising administering to a patient in need of such treatment an effective amount of a compound as claimed in claim 1, or a pharmaceutically acceptable salt thereof or solvate thereof.

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